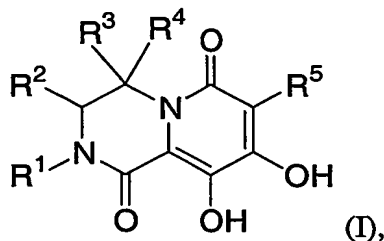


WHAT IS CLAIMED IS:

1. A compound of Formula (I), or a pharmaceutically acceptable salt thereof:



wherein

R¹ is C₁₋₆ alkyl which is substituted with 1 or 2 substituents each of which is independently:

- (1) C₃₋₈ cycloalkyl,
- (2) aryl,
- (3) a 5- or 6-membered saturated or mono-unsaturated heterocyclic ring containing from 1 to 4 heteroatoms independently selected from N, O and S,
- (4) a 5- or 6-membered heteroaromatic ring containing from 1 to 4 heteroatoms independently selected from N, O and S, or
- (5) a 9- or 10-membered fused bicyclic heterocycle containing from 1 to 4 heteroatoms independently selected from N, O and S, wherein at least one of the rings is aromatic;

wherein

- (A) each cycloalkyl is optionally substituted with from 1 to 3 substituents, each of which is independently halo, -C₁₋₆ alkyl, or -O-C₁₋₆ alkyl;
- (B) each aryl is optionally substituted with from 1 to 5 substituents each of which is independently

- (1) -C₁₋₆ alkyl, optionally substituted with from 1 to 3 substituents each of which is independently -OH, -O-C₁₋₆ alkyl, -O-C₁₋₆ haloalkyl, -CN, -NO₂, -N(R^aR^b), -C(=O)N(R^aR^b), -C(=O)R^a, -CO₂R^c, -S(O)_nR^c, -SO₂N(R^aR^b), -N(R^a)C(=O)R^b, -N(R^a)CO₂R^c, -N(R^a)SO₂R^c, -N(R^a)SO₂N(R^aR^b), -OC(=O)N(R^aR^b), or -N(R^a)C(=O)N(R^aR^b),

- 5 (2) -O-C₁₋₆ alkyl, optionally substituted with from 1 to 3 substituents each of which is independently -OH, -O-C₁₋₆ alkyl, -O-C₁₋₆ haloalkyl, -S(O)_nR^c, -C(=O)N(R^aR^b), -SO₂N(R^aR^b), -N(R^a)C(=O)R^b, -N(R^a)CO₂R^c, -N(R^a)SO₂R^c, -N(R^a)SO₂N(R^aR^b), -OC(=O)N(R^aR^b), or -N(R^a)C(=O)N(R^aR^b),
- 10 (3) -C₁₋₆ haloalkyl,
- (4) -O-C₁₋₆ haloalkyl,
- (5) -OH,
- (6) halo,
- (7) -CN,
- (8) -NO₂,
- (9) -N(R^aR^b),
- (10) -C(=O)N(R^aR^b),
- 15 (11) -C(=O)R^a,
- (12) -CO₂R^c,
- (13) -SR^c,
- (14) -S(=O)R^c,
- (15) -SO₂R^c,
- 20 (16) -N(R^a)SO₂R^c,
- (17) -SO₂N(R^aR^b),
- (18) -N(R^a)C(=O)R^b,
- (19) -N(R^a)CO₂R^c, or
- (20) phenyl;
- 25 (C) each saturated or mono-unsaturated heterocyclic ring is
- (i) optionally substituted with from 1 to 5 substituents each of which is independently halogen, -C₁₋₆ alkyl, -C₁₋₆ haloalkyl, -O-C₁₋₆ alkyl, -O-C₁₋₆ haloalkyl, or oxo; and
- 30 (ii) optionally substituted with 1 or 2 substituents each of which is independently aryl or a 5- or 6-membered heteroaromatic ring containing from 1 to 4 heteroatoms independently selected from N, O and S; and
- (D) each heteroaromatic ring or each fused bicyclic heterocycle is
- 35 (i) optionally substituted with from 1 to 7 substituents each of which is independently halogen, -C₁₋₆ alkyl, -C₁₋₆

haloalkyl, -O-C₁₋₆ alkyl, -O-C₁₋₆ haloalkyl, or oxo;

and

- (ii) optionally substituted with 1 or 2 substituents each of which is independently aryl or -C₁₋₆ alkyl-aryl;

5

R² is -H or -C₁₋₆ alkyl;

R³ is -H, -C₁₋₆ alkyl, -C₁₋₆ haloalkyl, or -C₁₋₆ alkyl substituted with one of -OH, -O-C₁₋₆ alkyl, -O-C₁₋₆ haloalkyl, -CN, -NO₂, -N(R^aR^b), -C(=O)N(R^aR^b),
 10 -C(=O)R^a, -CO₂R^c, -S(O)_nR^c, -SO₂N(R^aR^b), -N(R^a)C(=O)R^b, -N(R^a)CO₂R^c,
 -N(R^a)SO₂R^c, -N(R^a)SO₂N(R^aR^b), -OC(=O)N(R^aR^b), or -N(R^a)C(=O)N(R^aR^b);

R⁴ is:

- 15 (1) -H,
 (2) -C₁₋₆ alkyl optionally substituted with one of -OH, -O-C₁₋₆ alkyl, -O-C₁₋₆ haloalkyl, -CN, -NO₂, -N(R^aR^b), -C(=O)N(R^aR^b), -C(=O)R^a, -CO₂R^c, -S(O)_nR^c, -SO₂N(R^aR^b), -N(R^a)-C(R^b)=O, -N(R^a)SO₂R^c, -N(R^a)SO₂N(R^aR^b), -OC(=O)N(R^aR^b), -N(R^a)C(=O)N(R^aR^b), -O-C₁₋₆ alkyl-C(=O)N(R^aR^b), -S-C₁₋₆ alkyl-C(=O)N(R^aR^b), -N(R^a)-C₁₋₆ alkyl-C(=O)N(R^aR^b), or -N(SO₂R^c)-C₁₋₆ alkyl-C(=O)N(R^aR^b),
 20 (3) -C₁₋₆ haloalkyl,
 (4) -C(=O)R^a,
 (5) -CO₂R^c,
 25 (6) -C(=O)N(R^aR^b),
 (7) -SO₂N(R^aR^b),
 (8) -C₂₋₆ alkenyl,
 (9) -C₂₋₆ alkenyl-C(=O)-N(R^a)₂,
 (10) -C₂₋₅ alkynyl,
 30 (11) -C₂₋₅ alkynyl-CH₂N(R^a)₂,
 (12) -C₂₋₅ alkynyl-CH₂OR^a,
 (13) -C₂₋₅ alkynyl-CH₂S(O)_nR^c, or
 (14) -R^k,
 (15) -C₁₋₆ alkyl substituted with R^k,
 35 (16) -C₁₋₆ haloalkyl substituted with R^k,
 (17) -C₁₋₆ alkyl-O-R^k,

- (18) -C₁₋₆ alkyl-O-C₁₋₆ alkyl-R^k,
 (19) -C₁₋₆ alkyl-S(O)_n-R^k,
 (20) -C₁₋₆ alkyl-S(O)_n-C₁₋₆ alkyl-R^k,
 (21) -C₁₋₆ alkyl-N(R^a)-R^k,
 5 (22) -C₁₋₆ alkyl-N(R^a)-C₁₋₆ alkyl-R^k,
 (23) -C₁₋₆ alkyl-N(R^a)-C₁₋₆ alkyl-OR^k, with the proviso that the -N(R^a)-moiety and the -OR^k moiety are not both attached to the same carbon of the -C₁₋₆ alkyl- moiety,
 (24) -C₁₋₆ alkyl-C(=O)-R^k,
 10 (25) -C₁₋₆ alkyl-C(=O)N(R^a)-R^k,
 (26) -C₁₋₆ alkyl-N(R^a)C(=O)-R^k,
 (27) -C₁₋₆ alkyl-C(=O)N(R^a)-C₁₋₆ alkyl-R^k, or
 (28) -C₁₋₆ alkyl-N(R^a)-C₀₋₆ alkyl-S(O)_nR^k;
 wherein R^k is
 15 (i) aryl, which is optionally substituted with from 1 to 5 substituents each of which is independently -C₁₋₆ alkyl, -C₁₋₆ alkyl-OH, -C₁₋₆ alkyl-O-C₁₋₆ alkyl, -C₁₋₆ alkyl-O-C₁₋₆ haloalkyl, -C₁₋₆ alkyl-N(R^aR^b), -C₁₋₆ alkyl-C(=O)N(R^aR^b), -C₁₋₆ alkyl-C(=O)R^a, -C₁₋₆ alkyl-CO₂R^c, -C₁₋₆
 20 alkyl-S(O)_nR^c, -O-C₁₋₆ alkyl, -C₁₋₆ haloalkyl, -O-C₁₋₆ haloalkyl, -OH, halo, -N(R^aR^b), -C(=O)N(R^aR^b), -C(=O)R^a, -CO₂R^c, -S(O)_nR^c, or -SO₂N(R^aR^b);
 (ii) a 4- to 7-membered saturated or mono-unsaturated heterocyclic ring containing at least one carbon atom and from 1 to 4
 25 heteroatoms independently selected from N, O and S, wherein the heterocyclic ring is:
 (a) optionally substituted with from 1 to 5 substituents each of which is independently halogen, -C₁₋₆ alkyl, -C₁₋₆ haloalkyl, -O-C₁₋₆ alkyl, -O-C₁₋₆ haloalkyl, or oxo;
 30 and
 (b) optionally mono-substituted with aryl or HetA;
 wherein HetA is a 5- or 6-membered heteroaromatic ring containing from 1 to 4 heteroatoms independently selected from N, O and S, wherein the heteroaromatic ring is optionally fused with a benzene ring, and HetA is optionally substituted with from 1 to 4
 35

substituents each of which is independently -C₁₋₆ alkyl, -C₁₋₆ haloalkyl, -O-C₁₋₆ alkyl, -O-C₁₋₆ haloalkyl, or oxo; or

- (iii) a 5- or 6-membered heteroaromatic ring containing from 1 to 4 heteroatoms independently selected from N, O and S, wherein the heteroaromatic ring is optionally substituted with from optionally substituted with from 1 to 4 substituents each of which is independently -C₁₋₆ alkyl, -C₁₋₆ haloalkyl, -O-C₁₋₆ alkyl, -O-C₁₋₆ haloalkyl, or oxo;

or alternatively R³ and R⁴ are joined together to form C₅₋₈ cycloalkyl or a 5- to 7-membered saturated heterocyclic ring containing from 1 to 4 heteroatoms independently selected from N, O and S; wherein

the cycloalkyl is optionally substituted with from 1 to 3 substituents each of which is independently halo, -C₁₋₆ alkyl, or -O-C₁₋₆ alkyl; and

the heterocyclic ring is optionally substituted with from 1 to 4 substituents each of which is independently -C₁₋₆ alkyl, -C₁₋₆ haloalkyl, -O-C₁₋₆ alkyl, -O-C₁₋₆ haloalkyl, or oxo;

or alternatively:

(i) R² and R³ together form a direct bond to give a ring double bond, and R⁴ is an independent group as defined above;

(ii) R² and R³ together with the ring carbon atoms to which they are attached form a fused cyclopropyl ring which is optionally substituted at the non-fused cyclopropyl ring carbon with -OR^d, and R⁴ is -H; or

(iii) R² and R³ together with the ring carbon atoms to which they are attached form a fused phenyl ring or a fused pyridyl ring, and R⁴ is absent;

R⁵ is:

- (1) -H,
- (2) -C₁₋₆ alkyl,
- (3) -C₁₋₆ alkyl-N(R^aR^b),
- (4) -C₁₋₆ alkyl-C(=O)N(R^aR^b),
- (5) -C₁₋₆ alkyl-C(=O)R^a,
- (6) -C₁₋₆ alkyl-CO₂R^c,
- (7) -C₁₋₆ alkyl-SR^c,

- (8) -C₁₋₆ alkyl-S(=O)R^c,
 (9) -C₁₋₆ alkyl-SO₂R^c,
 (10) -C₁₋₆ alkyl-SO₂N(R^aR^b)
 (11) -C₁₋₆ haloalkyl,
 5 (12) -O-C₁₋₆ alkyl,
 (13) -O-C₁₋₆ haloalkyl,
 (14) halo,
 (15) -CN,
 (16) -C(=O)R^a,
 10 (17) -CO₂R^c,
 (18) -SR^c,
 (19) -S(=O)R^c,
 (20) -SO₂R^c,
 (21) -N(R^aR^b),
 15 (22) -C(=O)N(R^aR^b), or
 (23) -SO₂N(R^aR^b);
 (24) aryl
 (25) -C₁₋₆ alkyl-aryl
 (26) HetB,
 20 (27) -C₁₋₆ alkyl-HetB,
 (28) HetC, or
 (29) -C₁₋₆ alkyl-HetC,

wherein

25 HetB is a 5- or 6-membered saturated or mono-unsaturated ring
 containing from 1 to 4 heteroatoms independently selected from N, O
 and S, wherein the ring is optionally substituted with from 1 to 5
 substituents each of which is independently halogen, -C₁₋₆ alkyl,
 -C₁₋₆ haloalkyl, -O-C₁₋₆ alkyl, -O-C₁₋₆ haloalkyl, oxo, -C(=O)-C₁₋₆
 alkyl, -C(=O)-C₁₋₆ haloalkyl, or -C₁₋₆ alkyl-C₃₋₈ cycloalkyl; and

30 HetC is a 5- or 6-membered heteroaromatic ring containing
 from 1 to 4 heteroatoms independently selected from N, O and S,
 wherein the heteroaromatic ring is optionally substituted with from 1 to
 4 substituents each of which is independently -C₁₋₆ alkyl, -C₁₋₆
 haloalkyl, -O-C₁₋₆ alkyl, -O-C₁₋₆ haloalkyl, or oxo;

35 each R^a and R^b is independently -H or -C₁₋₆ alkyl;

each R^c is independently a -C₁₋₆ alkyl;

R^d is a -C₁₋₆ alkyl, allyl, or benzyl; and

5

each n is independently an integer equal to 0, 1 or 2.

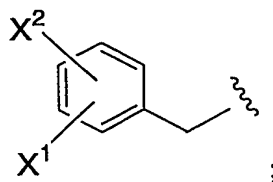
2. The compound according to claim 1, or a pharmaceutically acceptable salt thereof, wherein R¹ is -C₁₋₄ alkyl mono-substituted with aryl; wherein
10 the aryl is optionally substituted with from 1 to 4 substituents each of which is independently

- (1) -C₁₋₄ alkyl, optionally mono-substituted with -OH, -O-C₁₋₄ alkyl, -O-C₁₋₄ haloalkyl, -CN, -N(R^aR^b), -C(=O)N(R^aR^b), -C(=O)R^a, -CO₂R^c, -S(O)_nR^c, -SO₂N(R^aR^b), -N(R^a)C(=O)R^b, -N(R^a)CO₂R^c,
15 -N(R^a)SO₂R^c, -N(R^a)SO₂N(R^aR^b), -OC(=O)N(R^aR^b), or -N(R^a)C(=O)N(R^aR^b),
 - (2) -O-C₁₋₄ alkyl, optionally mono-substituted with -OH, -O-C₁₋₄ alkyl, -O-C₁₋₄ haloalkyl, -S(O)_nR^c, -N(R^a)-CO₂R^c, -C(=O)N(R^aR^b), -SO₂N(R^aR^b), -N(R^a)C(=O)R^b, -N(R^a)CO₂R^c, -N(R^a)SO₂R^c,
20 -N(R^a)SO₂N(R^aR^b), -OC(=O)N(R^aR^b), or -N(R^a)C(=O)N(R^aR^b),
 - (3) -C₁₋₄ haloalkyl,
 - (4) -O-C₁₋₄ haloalkyl,
 - (5) -OH,
 - (6) halo,
 - (7) -CN,
 - (8) -NO₂,
 - (9) -N(R^aR^b),
 - (10) -SR^c,
 - (11) -S(=O)R^c,
 - (12) -SO₂R^c,
 - (13) -N(R^a)SO₂R^c,
 - (14) -SO₂N(R^aR^b),
 - (15) -N(R^a)C(=O)R^b,
 - (16) -N(R^a)CO₂R^c, or
 - (17) phenyl.
- 25
30
35

3. The compound according to claim 2, or a pharmaceutically acceptable salt thereof, wherein R^1 is $-(CH_2)_{1-4}$ -phenyl, wherein the phenyl is optionally substituted with from 1 to 3 substituents each of which is independently

- (1) $-C_{1-4}$ alkyl, optionally mono-substituted with $-OH$, $-O-C_{1-4}$ alkyl, $-O-C_{1-4}$ haloalkyl, $-CN$, $-N(R^aR^b)$, $-C(=O)N(R^aR^b)$, $-C(=O)R^a$, $-CO_2R^c$, $-S(O)_nR^c$, or $-SO_2N(R^aR^b)$,
- (2) $-O-C_{1-4}$ alkyl,
- (3) $-C_{1-4}$ haloalkyl,
- (4) $-O-C_{1-4}$ haloalkyl,
- (5) $-OH$,
- (6) halo,
- (7) $-CN$,
- (8) $-NO_2$,
- (9) $-N(R^aR^b)$,
- (10) $-SR^c$,
- (11) $-S(=O)R^c$,
- (12) $-SO_2R^c$,
- (13) $-N(R^a)SO_2R^c$,
- (14) $-SO_2N(R^aR^b)$,
- (15) $-N(R^a)C(=O)R^b$,
- (16) $-N(R^a)CO_2R^c$, or
- (17) phenyl.

4. The compound according to claim 3, or a pharmaceutically acceptable salt thereof, wherein R^1 is:



wherein X^1 and X^2 are each independently

- (1) $-H$,
- (2) methyl,
- (3) ethyl,
- (4) methoxy,

- (5) ethoxy,
- (6) -CF₃,
- (7) fluoro,
- (8) bromo,
- 5 (9) chloro,
- (10) -CN,
- (11) -S-CH₃, or
- (12) phenyl.

10 5. The compound according to claim 4, or a pharmaceutically acceptable salt thereof, wherein R¹ is 4-fluorobenzyl.

6. The compound according to claim 1, or a pharmaceutically acceptable salt thereof, wherein

15

R² is -H or -C₁₋₄ alkyl;

R³ is -H or -C₁₋₄ alkyl; and

20 R⁴ is:

- (1) -H,
- (2) -C₁₋₄ alkyl optionally substituted with one of -OH, -O-C₁₋₄ alkyl, -O-C₁₋₄ haloalkyl, -CN, -N(R^aR^b), -C(=O)N(R^aR^b), -C(=O)R^a, -CO₂R^c, -S(O)_nR^c, -SO₂N(R^aR^b), -N(R^a)-C(R^b)=O, -N(R^a)SO₂R^b, or -N(R^a)SO₂N(R^aR^b),
- 25 (3) -C(=O)N(R^aR^b),
- (4) -R^k,
- (5) -C₁₋₄ alkyl substituted with R^k,
- (6) -C₁₋₄ alkyl-O-R^k, or
- 30 (7) -C₁₋₄ alkyl-O-C₁₋₄ alkyl-R^k.

7. The compound according to claim 1, or a pharmaceutically acceptable salt thereof, wherein R⁵ is:

- (1) -H,
- 35 (2) -C₁₋₄ alkyl,
- (3) -C₁₋₄ alkyl-N(R^aR^b),
- (4) -C₁₋₄ alkyl-C(=O)N(R^aR^b),

- (5) -C₁₋₄ alkyl-SO₂N(R^aR^b)
 (6) -C₁₋₄ haloalkyl,
 (7) halo,
 (8) -CN,
 5 (9) aryl
 (10) -C₁₋₄ alkyl-aryl
 (11) HetB,
 (12) -C₁₋₄ alkyl-HetB,
 (13) HetC, or
 10 (14) -C₁₋₄ alkyl-HetC,

wherein

15 HetB is a 5- or 6-membered saturated ring containing a total of from 1 to 4 heteroatoms independently selected from 1 to 4 N atoms, from 0 to 2 O atoms, and from 0 to 2 S atoms, wherein the saturated ring is optionally substituted with from 1 to 4 substituents each of which is independently halogen, -C₁₋₄ alkyl, -C₁₋₄ haloalkyl, -O-C₁₋₄ alkyl, -O-C₁₋₄ haloalkyl, oxo, -C(=O)-C₁₋₄ alkyl, -C(=O)-C₁₋₄ haloalkyl, or -C₁₋₄ alkyl-C₃₋₆ cycloalkyl; and

20 HetC is a 5- or 6-membered heteroaromatic ring containing a total of from 1 to 4 heteroatoms independently selected from 1 to 4 N atoms, from 0 to 2 O atoms, and from 0 to 2 S atoms, wherein the heteroaromatic ring is optionally substituted with from 1 to 3 substituents each of which is independently -C₁₋₄ alkyl, -C₁₋₄ haloalkyl, -O-C₁₋₄ alkyl, -O-C₁₋₄ haloalkyl, or oxo.

25

8. The compound according to claim 7, or a pharmaceutically acceptable salt thereof, wherein R⁵ is:

- (1) -H,
 (2) -C₁₋₄ alkyl,
 30 (3) -C₁₋₄ alkyl-N(R^aR^b),
 (4) halo,
 (5) -CN, or
 (6) -C₁₋₄ alkyl-HetB;

wherein

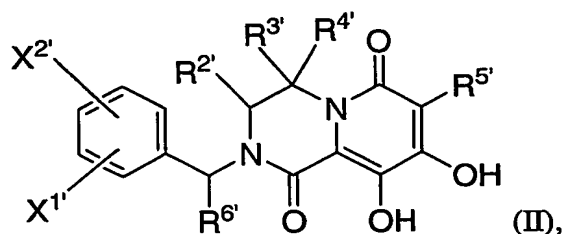
35

HetB is a 5- or 6-membered saturated ring containing 1 or 2 N atoms and carbon atoms, wherein the saturated ring is optionally

substituted with from 1 to 4 substituents each of which is independently halogen, -C₁₋₄ alkyl, -C₁₋₄ haloalkyl, -O-C₁₋₄ alkyl, -O-C₁₋₄ haloalkyl, oxo, -C(=O)-C₁₋₄ alkyl, -C(=O)-C₁₋₄ haloalkyl, or -C₁₋₄ alkyl-C₃₋₆ cycloalkyl.

5

9. A compound of Formula (II), or a pharmaceutically acceptable salt thereof:



wherein:

10

X^{1'} and X^{2'} are each independently:

- (1) -H,
- (2) C₁₋₄ alkyl,
- (2) -O-C₁₋₄ alkyl,
- 15 (3) -C₁₋₄ haloalkyl,
- (4) -O-C₁₋₄ haloalkyl,
- (5) halo,
- (6) -CN,
- (7) -S-C₁₋₄ alkyl, or
- 20 (8) phenyl;

R^{2'} is -H or -C₁₋₄ alkyl;

R^{3'} is -H or -C₁₋₄ alkyl;

25

R^{4'} is:

- (1) -H,
- (2) -C₁₋₄ alkyl optionally substituted with one of -OH, -N(R^{a'}R^{b'}), or -C(=O)N(R^{a'}R^{b'}),
- (3) -C(=O)N(R^{a'}R^{b'}),
- 30 (4) -(CH₂)₁₋₃-R^{k'},

(5) $-(\text{CH}_2)_{1-3}-\text{O}-\text{R}^{\text{k}'}$, or

(6) $-(\text{CH}_2)_{1-3}-\text{O}-(\text{CH}_2)_{1-3}-\text{R}^{\text{k}'}$;

wherein $\text{R}^{\text{k}'}$ is:

- 5 (i) phenyl, which is optionally substituted with from 1 to 3 substituents each of which is independently $-\text{C}_{1-4}$ alkyl, $-\text{O}-\text{C}_{1-4}$ alkyl, $-\text{C}_{1-4}$ haloalkyl, $-\text{O}-\text{C}_{1-4}$ haloalkyl, or halo; or
- 10 (ii) HetD, wherein HetD is a 5- or 6-membered saturated ring containing 1 or 2 N atoms, 0 or 1 S atoms, and a balance of carbon atoms, wherein the saturated ring is optionally substituted with from 1 to 4 substituents each of which is independently halo, $-\text{C}_{1-4}$ alkyl, $-\text{C}_{1-4}$ haloalkyl, $-\text{O}-\text{C}_{1-4}$ alkyl, $-\text{O}-\text{C}_{1-4}$ haloalkyl, or oxo;

or alternatively:

- 15 (i) $\text{R}^{2'}$ and $\text{R}^{3'}$ together form a direct bond to give a ring double bond, and $\text{R}^{4'}$ is an independent group as defined above;
- (ii) $\text{R}^{2'}$ and $\text{R}^{3'}$ together with the ring carbon atoms to which they are attached form a fused cyclopropyl ring which is optionally substituted at the non-fused cyclopropyl ring carbon with $-\text{OR}^{\text{d}'}$, and $\text{R}^{4'}$ is $-\text{H}$; or
- 20 (iii) $\text{R}^{2'}$ and $\text{R}^{3'}$ together with the ring carbon atoms to which they are attached form a fused phenyl ring or a fused pyridyl ring, and $\text{R}^{4'}$ is absent;

$\text{R}^{5'}$ is:

- 25 (1) $-\text{H}$,
- (2) $-\text{C}_{1-4}$ alkyl,
- (3) $-\text{C}_{1-4}$ alkyl- $\text{N}(\text{R}^{\text{a}'}\text{R}^{\text{b}'})$,
- (4) halo,
- (5) $-\text{CN}$, or
- (6) $-(\text{CH}_2)_{1-3}-\text{HetB}$;

30 wherein

HetB is a 5- or 6-membered saturated ring containing 1 or 2 N atoms, zero or 1 O atom, zero or 1 S atom, and a balance of carbon atoms, wherein the saturated ring is optionally substituted with from 1 to 4 substituents each of which is independently halogen, $-\text{C}_{1-4}$ alkyl, $-\text{C}_{1-4}$ haloalkyl, $-\text{O}-\text{C}_{1-4}$ alkyl, $-\text{O}-\text{C}_{1-4}$ haloalkyl, oxo, $-\text{C}(=\text{O})-\text{C}_{1-4}$ alkyl, $-\text{C}(=\text{O})-\text{C}_{1-4}$ haloalkyl, or $-\text{C}_{1-4}$ alkyl- C_{3-6} cycloalkyl;

35

R^{6'} is -H or methyl;

each R^{a'} and R^{b'} is independently -H or -C₁₋₄ alkyl; and

5

R^{d'} is -C₁₋₄ alkyl, allyl, or benzyl.

10. A compound according to claim 9, or a pharmaceutically acceptable salt thereof, wherein:

10

wherein X^{1'} and X^{2'} are each independently:

- (1) -H,
- (2) methyl,
- (2) -OCH₃,
- 15 (3) -CF₃,
- (4) -O-CF₃,
- (5) chloro,
- (6) fluoro,
- (7) bromo;
- 20 (6) -CN,
- (7) -S-CH₃, or
- (8) phenyl;

R^{2'} is -H or methyl;

25

R^{3'} is -H or methyl;

R^{4'} is:

- (1) -H,
- 30 (2) methyl,
- (3) -CH₂OH,
- (3) -C(=O)N(CH₃)₂,
- (4) -CH₂-R^{k'}, or
- (5) -CH₂-O-CH₂-R^{k'};
- 35 wherein R^{k'} is:

- (i) phenyl, which is optionally substituted with from 1 to 3 substituents each of which is independently -CH₃, -OCH₃, -CF₃, -OCF₃, chloro, bromo or fluoro; or
- (ii) HetD, wherein HetD is a 5- or 6-membered saturated ring containing 1 or 2 N atoms, 0 or 1 S atoms, and a balance of carbon atoms, wherein the saturated ring is optionally substituted with from 1 to 4 substituents each of which is independently chloro, bromo, fluoro, -CH₃, -CF₃, -OCH₃, -OCF₃, or oxo;

or alternatively:

- (i) R^{2'} and R^{3'} together form a direct bond to give a ring double bond, and R^{4'} is an independent group as defined above;
- (ii) R^{2'} and R^{3'} together with the ring carbon atoms to which they are attached form a fused cyclopropyl ring which is optionally substituted at the non-fused cyclopropyl ring carbon with -OMe, -OEt, -O-allyl, or -O-benzyl, and R^{4'} is -H; or
- (iii) R^{2'} and R^{3'} together with the ring carbon atoms to which they are attached form a fused phenyl ring or a fused pyridyl ring, and R^{4'} is absent;

R^{5'} is:

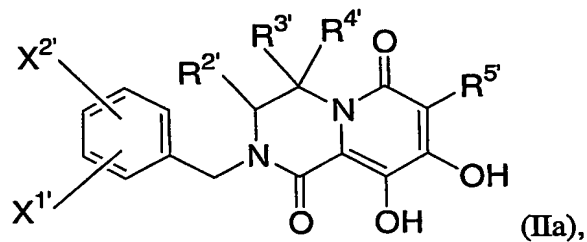
- (1) -H,
- (2) methyl,
- (3) -(CH₂)₁₋₂-N(CH₃)₂,
- (4) fluoro,
- (5) bromo,
- (6) iodo,
- (7) -CN, or
- (8) -CH₂-HetB;

wherein

HetB is a 5- or 6-membered saturated ring containing 1 or 2 N atoms, zero or 1 O atom, zero or 1 S atom, and a balance of carbon atoms, wherein the saturated ring is optionally substituted with from 1 to 4 substituents each of which is independently chloro, bromo, fluoro, -CH₃, -CF₃, -OCH₃, -OCF₃, oxo, -C(=O)-CH₃, -C(=O)-CF₃, or -CH₂-cyclopropyl; and

R^{6'} is -H or methyl.

11. The compound according to claim 9, which is a compound of Formula (IIa), or a pharmaceutically acceptable salt thereof:



wherein:

X^{1'} and X^{2'} are each independently:

- (1) -H,
- (2) C₁₋₄ alkyl,
- (2) -O-C₁₋₄ alkyl,
- (3) -C₁₋₄ haloalkyl,
- (4) -O-C₁₋₄ haloalkyl, or
- (5) halo;

R^{2'} is -H or -C₁₋₄ alkyl;

R^{3'} is -H or -C₁₋₄ alkyl;

or alternatively R^{2'} and R^{3'} together form a direct bond to give a ring double bond;

R^{4'} is:

- (1) -H,
- (2) -C₁₋₄ alkyl optionally substituted with one of -OH, -N(R^{a'}R^{b'}), or -C(=O)N(R^{a'}R^{b'}),
- (3) -C(=O)N(R^{a'}R^{b'}),
- (4) -(CH₂)₁₋₃-R^{k'},
- (5) -(CH₂)₁₋₃-O-R^{k'}, or
- (6) -(CH₂)₁₋₃-O-(CH₂)₁₋₃-R^{k'};

wherein R^{k'} is:

- (i) phenyl, which is optionally substituted with from 1 to 3 substituents each of which is independently -C₁₋₄ alkyl, -O-C₁₋₄ alkyl, -C₁₋₄ haloalkyl, -O-C₁₋₄ haloalkyl, or halo; or
- (ii) HetD, wherein HetD is a 5- or 6-membered saturated ring containing 1 or 2 N atoms, 0 or 1 S atoms, and a balance of carbon atoms, wherein the saturated ring is optionally substituted with from 1 to 4 substituents each of which is independently halo, -C₁₋₄ alkyl, -C₁₋₄ haloalkyl, -O-C₁₋₄ alkyl, -O-C₁₋₄ haloalkyl, or oxo;

R^{5'} is:

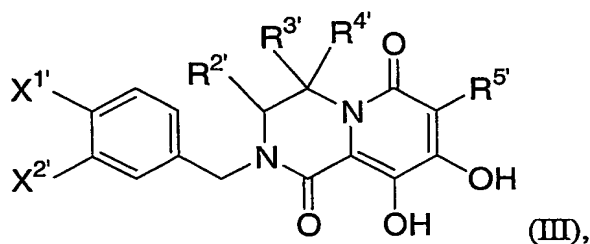
- (1) -H,
- (2) -C₁₋₄ alkyl,
- (3) -C₁₋₄ alkyl-N(R^{a'}R^{b'}),
- (4) halo,
- (5) -CN, or
- (6) -(CH₂)₁₋₃-HetB;

wherein

HetB is a 5- or 6-membered saturated ring containing 1 or 2 N atoms and carbon atoms, wherein the saturated ring is optionally substituted with from 1 to 4 substituents each of which is independently halogen, -C₁₋₄ alkyl, -C₁₋₄ haloalkyl, -O-C₁₋₄ alkyl, -O-C₁₋₄ haloalkyl, or oxo; and

each R^{a'} and R^{b'} is independently -H or -C₁₋₄ alkyl.

12. The compound according to claim 9, which is a compound of Formula (III), or a pharmaceutically acceptable salt thereof:



wherein:

X1' and X2' are each independently -H or halo.

13. The compound according to claim 12, or a pharmaceutically acceptable salt thereof,

wherein X1' and X2' are each independently -H, fluoro, chloro, or bromo;

R2' is -H or methyl;

R3' is -H or methyl;

R4' is:

- (1) -H,
- (2) methyl,
- (3) -CH₂OH,
- (3) -C(=O)N(CH₃)₂,
- (4) -CH₂-R^{k'}, or
- (5) -CH₂-O-CH₂-R^{k'};

wherein R^{k'} is:

- (i) phenyl, which is optionally substituted with from 1 to 3 substituents each of which is independently -CH₃, -OCH₃, -CF₃, -OCF₃, chloro, bromo or fluoro; or
- (ii) HetD, wherein HetD is a 5- or 6-membered saturated ring containing 1 or 2 N atoms, 0 or 1 S atoms, and a balance of carbon atoms, wherein the saturated ring is optionally substituted with from 1 to 4 substituents each of which is independently chloro, bromo, fluoro, -CH₃, -CF₃, -OCH₃, -OCF₃, or oxo; and

or alternatively:

- (i) R2' and R3' together form a direct bond to give a ring double bond, and R4' is an independent group as defined above;
- (ii) R2' and R3' together with the ring carbon atoms to which they are attached form a fused cyclopropyl ring which is optionally substituted at the non-fused cyclopropyl ring carbon with -ORD', and R4' is -H; or

(iii) R^{2'} and R^{3'} together with the ring carbon atoms to which they are attached form a fused phenyl ring or a fused pyridyl ring, and R^{4'} is absent;

R^{5'} is:

- 5 (1) -H,
- (2) methyl,
- (3) -(CH₂)₁₋₂-N(CH₃)₂,
- (4) fluoro,
- (5) bromo,
- 10 (6) iodo,
- (7) -CN, or
- (8) -CH₂-HetB;

wherein

15 HetB is a 5- or 6-membered saturated ring containing 1 or 2 N atoms, zero or 1 O atom, zero or 1 S atom, and a balance of carbon atoms, wherein the saturated ring is optionally substituted with from 1 to 4 substituents each of which is independently chloro, bromo, fluoro, -CH₃, -CF₃, -OCH₃, -OCF₃, oxo, -C(=O)-CH₃, -C(=O)-CF₃, or -CH₂-cyclopropyl.

20 14. The compound according to claim 13, or a pharmaceutically acceptable salt thereof, wherein X^{1'} is fluoro and X^{2'} is -H.

25 15. A compound selected from the group consisting of:

2-benzyl-8,9-dihydroxy-3,4-dihydro-2H-pyrido[1,2-*a*]pyrazine-1,6-dione;

2-(4-fluorobenzyl)-8,9-dihydroxy-3,4-dihydro-2H-pyrido[1,2-*a*]pyrazine-1,6-dione;

30 2-(4-fluorobenzyl)-8,9-dihydroxy-7-methyl-3,4-dihydro-2H-pyrido[1,2-*a*]pyrazine-1,6-dione;

2-(4-fluorobenzyl)-8,9-dihydroxy-7-bromo-3,4-dihydro-2H-pyrido[1,2-*a*]pyrazine-1,6-dione;

35

2-(4-fluorobenzyl)-8,9-dihydroxy-7-iodo-3,4-dihydro-2H-pyrido[1,2-*a*]pyrazine-1,6-dione;

2-(3-chlorobenzyl)-8,9-dihydroxy-3,4-dihydro-2H-pyrido[1,2-*a*]pyrazine-1,6-dione;

5

2-(4-chlorobenzyl)-8,9-dihydroxy-3,4-dihydro-2H-pyrido[1,2-*a*]pyrazine-1,6-dione

2-(3,4-dichlorobenzyl)-8,9-dihydroxy-3,4-dihydro-2H-pyrido[1,2-*a*]pyrazine-1,6-dione;

10

2-(3,4-difluorobenzyl)-8,9-dihydroxy-3,4-dihydro-2H-pyrido[1,2-*a*]pyrazine-1,6-dione

2-(3-chloro-4-fluorobenzyl)-8,9-dihydroxy-3,4-dihydro-2H-pyrido[1,2-*a*]pyrazine-1,6-dione

15

2-(4-fluorobenzyl)-8,9-dihydroxy-7-(piperidin-1-ylmethyl)-3,4-dihydro-2H-pyrido[1,2-*a*]pyrazine-1,6-dione;

20

2-(3-chloro-4-fluorobenzyl)-8,9-dihydroxy-7-(piperidin-1-ylmethyl)-3,4-dihydro-2H-pyrido[1,2-*a*]pyrazine-1,6-dione;

2-(4-fluorobenzyl)-8,9-dihydroxy-7-[(dimethylamino)methyl]-3,4-dihydro-2H-pyrido[1,2-*a*]pyrazine-1,6-dione;

25

2-(4-fluorobenzyl)-8,9-dihydroxy-2H-pyrido[1,2-*a*]pyrazine-1,6-dione

2-benzyl-8,9-dihydroxy-2H-pyrido[1,2-*a*]pyrazine-1,6-dione;

30

2-(4-fluorobenzyl)-8,9-dihydroxy-4-methyl-3,4-dihydro-2H-pyrido[1,2-*a*]pyrazine-1,6-dione

2-(4-fluorobenzyl)-8,9-dihydroxy-4,4-dimethyl-3,4-dihydro-2H-pyrido[1,2-*a*]pyrazine-1,6-dione;

35

2-(4-fluorobenzyl)-8,9-dihydroxy-3-methyl-3,4-dihydro-2H-pyrido[1,2-*a*]pyrazine-1,6-dione;

2-(4-fluorobenzyl)-8,9-dihydroxy-1,6-dioxo-1,3,4,6-tetrahydro-2H-pyrido[1,2-a]-pyrazine-7-carbonitrile;

- 5 2-(4-fluorobenzyl)-8,9-dihydroxy-7-[(4-methyl-3-oxopiperazin-1-yl)methyl]-3,4-dihydro-2H-pyrido[1,2-a]pyrazine-1,6-dione;

2-(4-fluorobenzyl)-8,9-dihydroxy-7-[(3-oxopiperazin-1-yl)methyl]-3,4-dihydro-2H-pyrido[1,2-a]pyrazine-1,6-dione;

10

4-[(benzyloxy)methyl]-2-(4-fluorobenzyl)-8,9-dihydroxy-3,4-dihydro-2H-pyrido[1,2-a]pyrazine-1,6-dione;

- 15 4-(hydroxymethyl)-2-(4-fluorobenzyl)-8,9-dihydroxy-3,4-dihydro-2H-pyrido[1,2-a]pyrazine-1,6-dione;

4-[(1,1-dioxido-1,2-thiazinan-2-yl)methyl]-2-(4-fluorobenzyl)-8,9-dihydroxy-3,4-dihydro-2H-pyrido[1,2-a]pyrazine-1,6-dione;

- 20 2-(4-fluorobenzyl)-8,9-dihydroxy-7-(piperidin-1-ylmethyl)-2H-pyrido[1,2-a]pyrazine-1,6-dione;

2-(4-fluorobenzyl)-8,9-dihydroxy-7-[(3-oxopiperazin-1-yl)methyl]-2H-pyrido[1,2-a]pyrazine-1,6-dione;

25

2-(4-fluorobenzyl)-8,9-dihydroxy-7-[(4-methyl-3-oxopiperazin-1-yl)methyl]-2H-pyrido[1,2-a]pyrazine-1,6-dione;

- 30 2-(4-fluorobenzyl)-8,9-dihydroxy-7-[(morpholin-4-yl)methyl]-2H-pyrido[1,2-a]pyrazine-1,6-dione;

2-(4-fluorobenzyl)-8,9-dihydroxy-7-[(thiomorpholin-4-yl)methyl]-2H-pyrido[1,2-a]pyrazine-1,6-dione;

- 35 2-[4-fluoro-2-(methylthio)benzyl]-8,9-dihydroxy-2H-pyrido[1,2-a]pyrazine-1,6-dione;

7-[(1-acetylpiperidin-4-yl)methyl]-2-(4-fluorobenzyl)-8,9-dihydroxy-3,4-dihydro-2H-pyrido[1,2-a]pyrazine-1,6-dione;

5 2-(4-fluorobenzyl)-8,9-dihydroxy-7-[[1-(trifluoroacetyl)piperidin-4-yl]methyl]-3,4-dihydro-2H-pyrido[1,2-a]pyrazine-1,6-dione;

7-[[1-(cyclopropylmethyl)piperidin-3-yl]methyl]-2-(4-fluorobenzyl)-8,9-dihydroxy-3,4-dihydro-2H-pyrido[1,2-a]pyrazine-1,6-dione;

10 7-[(1-acetylpiperidin-3-yl)methyl]-2-(4-fluorobenzyl)-8,9-dihydroxy-3,4-dihydro-2H-pyrido[1,2-a]pyrazine-1,6-dione;

7-[(1-acetylpiperidin-2-yl)methyl]-2-(4-fluorobenzyl)-8,9-dihydroxy-3,4-dihydro-2H-pyrido[1,2-a]pyrazine-1,6-dione;

15

7-[[1-(cyclopropylmethyl)piperidin-2-yl]methyl]-2-(4-fluorobenzyl)-8,9-dihydroxy-3,4-dihydro-2H-pyrido[1,2-a]pyrazine-1,6-dione;

2-(3-cyanobenzyl)-8,9-dihydroxy-3,4-dihydro-2H-pyrido[1,2-a]pyrazine-1,6-dione

20

2-(biphenyl-3-ylmethyl)-8,9-dihydroxy-3,4-dihydro-2H-pyrido[1,2-a]pyrazine-1,6-dione

(±)-1-[(benzyloxy)methyl]-2-(4-fluorobenzyl)-4,5-dihydroxy-1,1a,2,8a-tetrahydrocyclo-propa[e]pyrido[1,2-a]pyrazine-3,7-dione;

25

(±)-1-(methoxymethyl)-2-(4-fluorobenzyl)-4,5-dihydroxy-1,1a,2,8a-tetrahydrocyclo-propa[e]pyrido[1,2-a]pyrazine-3,7-dione;

30 (±)-1-[(allyloxy)methyl]-2-(4-fluorobenzyl)-4,5-dihydroxy-1,1a,2,8a-tetrahydrocyclo-propa[e]pyrido[1,2-a]pyrazine-3,7-dione;

(±)-1-(ethoxymethyl)-2-(4-fluorobenzyl)-4,5-dihydroxy-1,1a,2,8a-tetrahydrocyclo-propa[e]pyrido[1,2-a]pyrazine-3,7-dione;

35

(±)-1-(n-propoxymethyl)-2-(4-fluorobenzyl)-4,5-dihydroxy-1,1a,2,8a-tetrahydrocyclo-propa[e]pyrido[1,2-a]pyrazine-3,7-dione;

2-[1-(4-fluorophenyl)ethyl]-8,9-dihydroxy-3,4-dihydro-2H-pyrido[1,2-*a*]pyrazine-1,6-dione;

5 5-(4-fluorobenzyl)-7,8-dihydroxy-5H-pyrido[1,2-*a*]quinoxaline-6,10-dione;

5-(4-fluorobenzyl)-7,8-dihydroxy-5H-pyrido[1,2-*a*:3',2'-*e*]pyrazine-6,10-dione;

5-(4-fluorobenzyl)-7,8-dihydroxy-5H-pyrido[1,2-*a*:2',3'-*e*]pyrazine-6,10-dione;

10

and pharmaceutically acceptable salts thereof.

16. A pharmaceutical composition comprising a therapeutically effective amount of a compound according to claim 1, or a pharmaceutically acceptable salt thereof, and a pharmaceutically acceptable carrier.

15

17. A method of inhibiting HIV integrase in a subject in need thereof which comprises administering to the subject a therapeutically effective amount of the compound according to claim 1, or a pharmaceutically acceptable salt thereof.

20

18. A method for preventing or treating infection by HIV or for preventing, treating or delaying the onset of AIDS in a subject in need thereof which comprises administering to the subject a therapeutically effective amount of the compound according to claim 1, or a pharmaceutically acceptable salt thereof.

25

19. A pharmaceutical composition which comprises the product prepared by combining an effective amount of a compound according to claim 1, or a pharmaceutically acceptable salt thereof, and a pharmaceutically acceptable carrier.

30

20. A combination useful for inhibiting HIV integrase, for treating or preventing infection by HIV, or for preventing, treating or delaying the onset of AIDS, which is a therapeutically effective amount of a compound according to claim 1, or a pharmaceutically acceptable salt thereof, and a therapeutically effective amount of an HIV infection/AIDS antiviral agent selected from the group consisting of HIV protease inhibitors, non-nucleoside HIV reverse transcriptase inhibitors and nucleoside HIV reverse transcriptase inhibitors.

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